STIC-ILL

From:

Prouty, Rebecca

Sent:

Monday, February 11, 2002 2:18 PM

To: Subject:

STIC-ILL **ILL Request**

Art Unit 1652 10A13, 308-4000 Mailbox: 10C01

Serial Number: 09/538,248

Please provide the following reference(s):

L169 ANSWER 7 OF 53 MEDLINE **DUPLICATE 7**

TI c-Src mediates mitogenic signals and associates with cytoskeletal proteins upon vascular endothelial growth factor stimulation in Kaposi's sarcoma cells.

SO JOURNAL OF IMMUNOLOGY, (2000 Feb 1) 164 (3) 1169-74. Journal code: IFB; 2985117R. ISSN: 0022-1767.

AU Munshi N; Groopman J E; Gill P S; Ganju R K

AN 2000109065 MEDLINE 0/1301.F33

±169 ANSWER 11 OF 53 SCISEARCH COPYRIGHT 2002 ISI (R) DUPLICATE 10

TI SRC-family kinase antagonist (PP2) inhibits VEGF-stimulated

VE-cadherin tyrosine phosphorylation in microvascular endothelial cells.

SO FASEB JOURNAL, (15 MAR 2000) Vol. 14, No. 4, pp. A145-A145. Publisher: FEDERATION AMER SOC EXP BIOL, 9650 ROCKVILLE PIKE, BETHESDA, MD

20814-3998. ISSN: 0892-6638. AU Cooke L S (Reprint); Forough R; Dawson N; Parrish A; Hoffman P; Kilgannon P; Granger H G

AN 2000:328497 SCISEARCH

L169 ANSWER 15 OF 53 MEDLINE **DUPLICATE 12**

TI Vascular endothelial growth factor signals endothelial cell production of nitric oxide and prostacyclin through flk-1/KDR activation of c-Src.

SO JOURNAL OF BIOLÓGICAL CHEMISTRY, (1999 Aug 27) 274 (35) 25130-5.

Journal code: HIV; 2985121R. ISSN: 0021-9258.

AU He H; Venema V J; Gu X; Venema R C; Marrero M B; Caldwell R B

AN 1999387002 MEDLINE

L169 ANSWER 30 OF 53 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.DUPLICATE 21

TI Tyrosine kinases in disease: Overview of kinase inhibitors as therapeutic agents and current drugs in clinical trials.

SO Expert Opinion on Investigational Drugs, (1998) 7/4 (553-573).

Refs: 190 ISSN: 1354-3784 CODEN: EOIDER

AU Strawn L.M.; Shawver L.K.

AN 1998120483 EMBASE

L169 ANSWER 52 OF 53 MEDLINE **DUPLICATE 35**

TI Different signal transduction properties of KDR and Flt1, two receptors for vascular endothelial growth factor.

SO JOURNAL OF BIOLOGICAL CHEMISTRY, (1994 Oct 28) 269 (43) 26988-95.

Journal code: HIV; 2985121R. ISSN: 0021-9258. AU Waltenberger J; Claesson-Welsh L; Siegbahn A; Shibuya M; Heldin CH

AN 95014567 MEDLINE

NO (wrong page page)
AS 2/14

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AU Waltenberger J; Claesson-Welsh L; Siegbahn A; Shibuya M; Heldin C H

AN 95014567 MEDLINE

```
* * * * * * * * * * * * * * STN Columbus
=> fil reg
=> s 172889-26-8 or 172889-26-9
             1 172889-26-8
                  (172889-26-8/RN)
              0 172889-26-9
                  (172889-26-9/RN)
             1 172889-26-8 OR 172889-26-9
L2
\Rightarrow s 172889-26-8 or 172889-27-9
             1 172889-26-8
                  (172889-26-8/RN)
              1 172889-27-9
                  (172889-27-9/RN)
              2 172889-26-8 OR 172889-27-9
L3
=> d tot
L3
     ANSWER 1 OF 2 REGISTRY COPYRIGHT 2002 ACS
RN
     172889-27-9 REGISTRY
     1H-Pyrazolo[3,4-d]pyrimidin-4-amine, 3-(4-chlorophenyl)-1-(1,1-
CN
     dimethylethyl) - (9CI) (CA INDEX NAME)
OTHER NAMES:
     PP 2
CN
     3D CONCORD
FS
     C15 H16 Cl N5
MF
SR
                   CA, CAPLUS, CHEMCATS, TOXCENTER, TOXLIT, USPATFULL
LC
     STN Files:
       t-Bu
N.
   NH<sub>2</sub>
**PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT**
               10 REFERENCES IN FILE CA (1967 TO DATE)
               2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
               10 REFERENCES IN FILE CAPLUS (1967 TO DATE)
     ANSWER 2 OF 2 REGISTRY COPYRIGHT 2002 ACS
L3
     172889-26-8 REGISTRY
RN
     1H-Pyrazolo[3,4-d]pyrimidin-4-amine, 1-(1,1-dimethylethyl)-3-(4-dimethylethyl)
CN
     methylphenyl) - (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     PP 1
     3D CONCORD
FS
     C16 H19 N5
MF
```

SR CA LC STN Files: CA, CAPLUS, CHEMCATS, TOXCENTER, TOXLIT, USPATFULL

t-Bu

NH₂

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 20 REFERENCES IN FILE CA (1967 TO DATE)
- 1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 20 REFERENCES IN FILE CAPLUS (1967 TO DATE)

=> log ySINCE FILE TOTAL COST IN U.S. DOLLARS SESSION ENTRY 4.76 35.36 FULL ESTIMATED COST TOTAL SINCE FILE DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) ENTRY SESSION 0.00 -0.62 CA SUBSCRIBER PRICE

=> fil .bec
COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
0.15

FILES 'MEDLINE, SCISEARCH, LIFESCI, BIOTECHDS, BIOSIS, EMBASE, HCAPLUS, NTIS, ESBIOBASE, BIOTECHNO, WPIDS' ENTERED AT 11:48:22 ON 11 FEB 2002 ALL COPYRIGHTS AND RESTRICTIONS APPLY. SEE HELP USAGETERMS FOR DETAILS.

11 FILES IN THE FILE LIST

=> s vascula?(3a)(permeab? or leak?)

FILE 'MEDLINE'

298739 VASCULA?

86873 PERMEAB?

31878 LEAK?

L1 10344 VASCULA?(3A)(PERMEAB? OR LEAK?)

FILE 'SCISEARCH'

174301 VASCULA?

80092 PERMEAB?

38371 LEAK?

L2 6521 VASCULA?(3A) (PERMEAB? OR LEAK?)

FILE 'LIFESCI'

20055 VASCULA?

```
15152 PERMEAB?
          4677 LEAK?
          1032 VASCULA? (3A) (PERMEAB? OR LEAK?)
L3
FILE 'BIOTECHDS'
          1477 VASCULA?
          2429 PERMEAB?
           751 LEAK?
            46 VASCULA? (3A) (PERMEAB? OR LEAK?)
T.4
FILE 'BIOSIS'
        812639 VASCULA?
         81574 PERMEAB?
         29396 LEAK?
         8035 VASCULA?(3A) (PERMEAB? OR LEAK?)
FILE 'EMBASE'
        315413 VASCULA?
         69207 PERMEAB?
         29788 LEAK?
          5851 VASCULA? (3A) (PERMEAB? OR LEAK?)
L6
FILE 'HCAPLUS'
        109683 VASCULA?
        174309 PERMEAB?
         65622 LEAK?
L7
          5809 VASCULA?(3A)(PERMEAB? OR LEAK?)
FILE 'NTIS'
          2276 VASCULA?
         12462 PERMEAB?
         14526 LEAK?
            79 VASCULA? (3A) (PERMEAB? OR LEAK?)
L8
FILE 'ESBIOBASE'
         45773 VASCULA?
         38894 PERMEAB?
          6664 LEAK?
          1555 VASCULA? (3A) (PERMEAB? OR LEAK?)
L9
FILE 'BIOTECHNO'
         26349 VASCULA?
         15091 PERMEAB?
          3867 LEAK?
          1053 VASCULA? (3A) (PERMEAB? OR LEAK?)
L10
FILE 'WPIDS'
         13282 VASCULA?
         89571 PERMEAB?
        107887 LEAK?
           300 VASCULA? (3A) (PERMEAB? OR LEAK?)
L11
TOTAL FOR ALL FILES
        40625 VASCULA?(3A)(PERMEAB? OR LEAK?)
=> s 112(8a)(inhibit? or decreas?)
FILE 'MEDLINE'
        948649 INHIBIT?
        760115 DECREAS?
```

٠.

```
663 L1 (8A) (INHIBIT? OR DECREAS?)
L13
FILE 'SCISEARCH'
        732043 INHIBIT?
        691479 DECREAS?
           448 L2 (8A) (INHIBIT? OR DECREAS?)
L14
FILE 'LIFESCI'
        262888 INHIBIT?
        182622 DECREAS?
            90 L3 (8A) (INHIBIT? OR DECREAS?)
L15
FILE 'BIOTECHDS'
         33375 INHIBIT?
         15496 DECREAS?
             6 L4 (8A) (INHIBIT? OR DECREAS?)
L16
FILE 'BIOSIS'
       1022135 INHIBIT?
        885276 DECREAS?
L17
           727 L5 (8A) (INHIBIT? OR DECREAS?)
FILE 'EMBASE'
        843314 INHIBIT?
        732340 DECREAS?
           672 L6 (8A) (INHIBIT? OR DECREAS?)
L18
FILE 'HCAPLUS'
       1461279 INHIBIT?
       1871874 DECREAS?
           880 L7 (8A) (INHIBIT? OR DECREAS?)
L19
FILE 'NTIS'
         19267 INHIBIT?
         49407 DECREAS?
             4 L8 (8A) (INHIBIT? OR DECREAS?)
L20
FILE 'ESBIOBASE'
        275631 INHIBIT?
        217812 DECREAS?
           172 L9 (8A) (INHIBIT? OR DECREAS?)
L21
FILE 'BIOTECHNO'
        252731 INHIBIT?
        144633 DECREAS?
            76 L10(8A)(INHIBIT? OR DECREAS?)
L22
FILE 'WPIDS'
        172552 INHIBIT?
        163615 DECREAS?
            57 L11(8A) (INHIBIT? OR DECREAS?)
L23
TOTAL FOR ALL FILES
          3795 L12(8A)(INHIBIT? OR DECREAS?)
=> s 124 and (pyrazolopyrimidine# or ppl or pp2 or src or yes or 172889-26-8 or
172889-27-9)
FILE 'MEDLINE'
            64 PYRAZOLOPYRIMIDINE#
```

```
973 PP1
           163 PP2
         10746 SRC
          2769 YES
             0 172889-26-8
             0 172889-27-9
             1 L13 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L25
               172889-26-8 OR 172889-27-9)
FILE 'SCISEARCH'
            99 PYRAZOLOPYRIMIDINE#
           979 PP1
           182 PP2
         10238 SRC
          2485 YES
             0 172889-26-8
             0 172889-27-9
             1 L14 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L26
               172889-26-8 OR 172889-27-9)
FILE 'LIFESCI'
            27 PYRAZOLOPYRIMIDINE#
           360 PP1
            43 PP2
          4432 SRC
           373 YES
             0 172889-26-8
             0 172889-27-9
             O L15 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L27
               172889-26-8 OR 172889-27-9)
FILE 'BIOTECHDS'
             2 PYRAZOLOPYRIMIDINE#
            27 PP1
            11 PP2
           135 SRC
            24 YES
             0 172889-26-8
             0 172889-27-9
             O L16 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L28
               172889-26-8 OR 172889-27-9)
FILE 'BIOSIS'
           115 PYRAZOLOPYRIMIDINE#
          1104 PP1
           228 PP2
         10426 SRC
          1520 YES
             0 172889-26-8
             0 172889-27-9
             2 L17 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L29
               172889-26-8 OR 172889-27-9)
FILE 'EMBASE'
           122 PYRAZOLOPYRIMIDINE#
           893 PP1
           123 PP2
          7957 SRC
          1922 YES
```

```
0 172889-26-8
             0 172889-27-9
             1 L18 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L30
               172889-26-8 OR 172889-27-9)
FILE 'HCAPLUS'
          1137 PYRAZOLOPYRIMIDINE#
          1187 PP1
           274 PP2
         10766 SRC
          1478 YES
            20 172889-26-8
            10 172889-27-9
             2 L19 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L31
               172889-26-8 OR 172889-27-9)
FILE 'NTIS'
             1 PYRAZOLOPYRIMIDINE#
            15 PP1
             2 PP2
          1937 SRC
           307 YES
             0 172889-26-8
             0 172889-27-9
             O L20 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L32
               172889-26-8 OR 172889-27-9)
FILE 'ESBIOBASE'
             9 PYRAZOLOPYRIMIDINE#
           694 PP1
           109 PP2
          4886 SRC
           463 YES
             0 172889
         44754 26
        237842 8
             0 172889-26-8
                 (172889(W)26(W)8)
             0 172889
         42591 27
        177442 9
             0 172889-27-9
                 (172889(W)27(W)9)
             1 L21 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L33
               172889-26-8 OR 172889-27-9)
FILE 'BIOTECHNO'
            15 PYRAZOLOPYRIMIDINE#
           571 PP1
            76 PP2
          5674 SRC
           307 YES
             0 172889-26-8
             0 172889-27-9
             1 L22 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L34
                172889-26-8 OR 172889-27-9)
```

FILE 'WPIDS'

71 PYRAZOLOPYRIMIDINE#

```
140 PP1
            69 PP2
           462 SRC
           676 YES
             0 172889
        300469 26
       1687338 8
             0 172889-26-8
                 (172889(W)26(W)8)
             0 172889
        186349 27
       1310082 9
             0 172889-27-9
                 (172889(W)27(W)9)
             3 L23 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L35
               172889-26-8 OR 172889-27-9)
TOTAL FOR ALL FILES
            12 L24 AND (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR
L36
               172889-26-8 OR 172889-27-9)
=> s vascular endothelial or vpf or vegf
FILE 'MEDLINE'
        279253 VASCULAR
         78248 ENDOTHELIAL
         10945 VASCULAR ENDOTHELIAL
                 (VASCULAR (W) ENDOTHELIAL)
           278 VPF
          4522 VEGF
         11478 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L37
FILE 'SCISEARCH'
        160520 VASCULAR
        101259 ENDOTHELIAL
         13251 VASCULAR ENDOTHELIAL
                 (VASCULAR (W) ENDOTHELIAL)
           387 VPF
          6026 VEGF
         14872 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L38
FILE 'LIFESCI'
         18149 "VASCULAR"
         13662 "ENDOTHELIAL"
          2261 VASCULAR ENDOTHELIAL
                 ("VASCULAR"(W)"ENDOTHELIAL")
            70 VPF
           918 VEGF
          2314 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L39
FILE 'BIOTECHDS'
          1340 VASCULAR
          1266 ENDOTHELIAL
           364 VASCULAR ENDOTHELIAL
                 (VASCULAR (W) ENDOTHELIAL)
            11 VPF
           159 VEGF
           377 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L40
```

FILE 'BIOSIS'

```
793732 VASCULAR
        101898 ENDOTHELIAL
         16300 VASCULAR ENDOTHELIAL
                 (VASCULAR (W) ENDOTHELIAL)
           342 VPF
          6383 VEGF
         16826 VASCULAR ENDOTHELIAL OR VPF OR VEGF
1.41
FILE 'EMBASE'
        294094 "VASCULAR"
         74125 "ENDOTHELIAL"
         10117 VASCULAR ENDOTHELIAL
                 ("VASCULAR"(W) "ENDOTHELIAL")
           269 VPF
          4223 VEGF
         10689 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L42
FILE 'HCAPLUS'
        102821 VASCULAR
         61437 ENDOTHELIAL
         12482 VASCULAR ENDOTHELIAL
                 (VASCULAR(W)ENDOTHELIAL)
           311 VPF
          4984 VEGF
         12861 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L43
FILE 'NTIS'
          2097 VASCULAR
           579 ENDOTHELIAL
            54 VASCULAR ENDOTHELIAL
                 (VASCULAR(W) ENDOTHELIAL)
            36 VPF
            25 VEGF
           101 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L44
FILE 'ESBIOBASE'
         42688 VASCULAR
         36302 ENDOTHELIAL
          5684 VASCULAR ENDOTHELIAL
                 (VASCULAR(W) ENDOTHELIAL)
           161 VPF
          3094 VEGF
          6043 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L45
FILE 'BIOTECHNO'
         24329 VASCULAR
         21443 ENDOTHELIAL
          4615 VASCULAR ENDOTHELIAL
                 (VASCULAR (W) ENDOTHELIAL)
           157 VPF
          2498 VEGF
          4902 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L46
FILE 'WPIDS'
         12176 VASCULAR
          2932 ENDOTHELIAL
           831 VASCULAR ENDOTHELIAL
                  (VASCULAR (W) ENDOTHELIAL)
            50 VPF
```

```
533 VEGF
           980 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L47
TOTAL FOR ALL FILES
         81443 VASCULAR ENDOTHELIAL OR VPF OR VEGF
L48
=> s (pyrazolopyrimidine# or pp1 or pp2 or src or yes or 172889-26-8 or 172889-27-9)
FILE 'MEDLINE'
            64 PYRAZOLOPYRIMIDINE#
           973 PP1
           163 PP2
         10746 SRC
          2769 YES
             0 172889-26-8
             0 172889-27-9
         14100 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L49
               OR 172889-27-9)
FILE 'SCISEARCH'
            99 PYRAZOLOPYRIMIDINE#
           979 PP1
           182 PP2
         10238 SRC
          2485 YES
             0 172889-26-8
             0 172889-27-9
         13461 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L50
               OR 172889-27-9)
FILE 'LIFESCI'
            27 PYRAZOLOPYRIMIDINE#
           360 PP1
            43 PP2
          4432 SRC
           373 YES
             0 172889-26-8
             0 172889-27-9
          5003 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L51
               OR 172889-27-9)
FILE 'BIOTECHDS'
             2 PYRAZOLOPYRIMIDINE#
            27 PP1
            11 PP2
           135 SRC
            24 YES
             0 172889-26-8
             0 172889-27-9
           188 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L52
               OR 172889-27-9)
FILE 'BIOSIS'
           115 PYRAZOLOPYRIMIDINE#
          1104 PP1
           228 PP2
         10426 SRC
          1520 YES
             0 172889-26-8
             0 172889-27-9
```

```
12726 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L53
              OR 172889-27-9)
FILE 'EMBASE'
           122 PYRAZOLOPYRIMIDINE#
           893 PP1
           123 PP2
          7957 SRC
          1922 YES
             0 172889-26-8
             0 172889-27-9
         10540 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L54
               OR 172889-27-9)
FILE 'HCAPLUS'
          1137 PYRAZOLOPYRIMIDINE#
          1187 PP1
           274 PP2
         10766 SRC
          1478 YES
            20 172889-26-8
            10 172889-27-9
         14195 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L55
               OR 172889-27-9)
FILE 'NTIS'
             1 PYRAZOLOPYRIMIDINE#
            15 PP1
             2 PP2
          1937 SRC
           307 YES
             0 172889-26-8
             0 172889-27-9
          2260 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L56
               OR 172889-27-9)
FILE 'ESBIOBASE'
             9 PYRAZOLOPYRIMIDINE#
           694 PP1
           109 PP2
          4886 SRC
           463 YES
             0 172889
         44754 26
        237842 8
             0 172889-26-8
                 (172889(W)26(W)8)
             0 172889
         42591 27
        177442 9
             0 172889-27-9
                 (172889(W)27(W)9)
          5828 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L57
               OR 172889-27-9)
FILE 'BIOTECHNO'
            15 PYRAZOLOPYRIMIDINE#
           571 PP1
```

76 PP2

```
5674 SRC
          307 YES
            0 172889-26-8
            0 172889-27-9
         6329 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L58
              OR 172889-27-9)
FILE 'WPIDS'
           71 PYRAZOLOPYRIMIDINE#
          140 PP1
           69 PP2
           462 SRC
           676 YES
            0 172889
       300469 26
       1687338 8
             0 172889-26-8
                (172889(W)26(W)8)
            0 172889
       186349 27
       1310082 9
             0 172889-27-9
                (172889(W)27(W)9)
          1322 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L59
               OR 172889-27-9)
TOTAL FOR ALL FILES
         85952 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR SRC OR YES OR 172889-26-8
L60
              OR 172889-27-9)
=> s 148 and 160
FILE 'MEDLINE'
           90 L37 AND L49
L61
FILE 'SCISEARCH'
          96 L38 AND L50
L62
FILE 'LIFESCI'
      21 L39 AND L51
L63
FILE 'BIOTECHDS'
L64
           0 L40 AND L52
FILE 'BIOSIS'
      97 L41 AND L53
L65
FILE 'EMBASE'
L66 68 L42 AND L54
FILE 'HCAPLUS'
         109 L43 AND L55
L67
FILE 'NTIS'
           0 L44 AND L56
L68
FILE 'ESBIOBASE'
           56 L45 AND L57
```

FILE 'BIOTECHNO'

```
43 L46 AND L58
L70
FILE 'WPIDS'
         16 L47 AND L59
TOTAL FOR ALL FILES
         596 L48 AND L60
L72
=> s 148(15a)160
FILE 'MEDLINE'
     40 L37(15A)L49
L73
FILE 'SCISEARCH'
          44 L38(15A)L50
L74
FILE 'LIFESCI'
L75
    17 L39(15A)L51
FILE 'BIOTECHDS'
     0 L40(15A)L52
L76
FILE 'BIOSIS'
L77 44 L41(15A)L53
FILE 'EMBASE'
          39 L42(15A)L54
FILE 'HCAPLUS'
          43 L43(15A)L55
FILE 'NTIS'
           0 L44(15A)L56
L80
FILE 'ESBIOBASE'
     32 L45(15A)L57
L81
FILE 'BIOTECHNO'
         27 L46(15A)L58
L82
FILE 'WPIDS'
     6 L47(15A)L59
L83
TOTAL FOR ALL FILES
L84 292 L48(15A) L60
=> s (src or yes)(8a)(inhibit? or decreas?)
FILE 'MEDLINE'
        10746 SRC
         2769 YES
       948649 INHIBIT?
       760115 DECREAS?
        1260 (SRC OR YES)(8A)(INHIBIT? OR DECREAS?)
L85
FILE 'SCISEARCH'
        10238 SRC
         2485 YES
```

732043 INHIBIT? 691479 DECREAS?

```
1225 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L86
FILE 'LIFESCI'
          4432 SRC
           373 YES
        262888 INHIBIT?
        182622 DECREAS?
           494 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L87
FILE 'BIOTECHDS'
           135 SRC
            24 YES
         33375 INHIBIT?
         15496 DECREAS?
            13 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L88
FILE 'BIOSIS'
         10426 SRC
          1520 YES
       1022135 INHIBIT?
        885276 DECREAS?
          1408 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L89
FILE 'EMBASE'
          7957 SRC
          1922 YES
        843314 INHIBIT?
        732340 DECREAS?
          1162 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L90
FILE 'HCAPLUS'
         10766 SRC
          1478 YES
       1461279 INHIBIT?
       1871874 DECREAS?
          1416 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L91
FILE 'NTIS'
          1937 SRC
           307 YES
         19267 INHIBIT?
         49407 DECREAS?
            22 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L92
FILE 'ESBIOBASE'
          4886 SRC
           463 YES
        275631 INHIBIT?
        217812 DECREAS?
           821 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L93
FILE 'BIOTECHNO'
          5674 SRC
           307 YES
        252731 INHIBIT?
        144633 DECREAS?
           729 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L94
```

FILE 'WPIDS'

```
462 SRC
           676 YES
        172552 INHIBIT?
        163615 DECREAS?
            80 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L95
TOTAL FOR ALL FILES
          8630 (SRC OR YES) (8A) (INHIBIT? OR DECREAS?)
L96
=> s (pyrazolopyrimidine# or pp1 or pp2 or 172889-26-8 or 172889-27-9)
FILE 'MEDLINE'
            64 PYRAZOLOPYRIMIDINE#
           973 PP1
           163 PP2
             0 172889-26-8
             0 172889-27-9
          1153 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L97
FILE 'SCISEARCH'
            99 PYRAZOLOPYRIMIDINE#
           979 PP1
           182 PP2
             0 172889-26-8
             0 172889-27-9
          1216 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L98
FILE 'LIFESCI'
            27 PYRAZOLOPYRIMIDINE#
           360 PP1
            43 PP2
             0 172889-26-8
             0 172889-27-9
           416 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L99
FILE 'BIOTECHDS'
             2 PYRAZOLOPYRIMIDINE#
            27 PP1
            11 PP2
             0 172889-26-8
             0 172889-27-9
            35 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L100
FILE 'BIOSIS'
           115 PYRAZOLOPYRIMIDINE#
          1104 PP1
           228 PP2
             0 172889-26-8
             0 172889-27-9
          1385 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L101
               )
FILE 'EMBASE'
           122 PYRAZOLOPYRIMIDINE#
           893 PP1
           123 PP2
```

```
0 172889-26-8
             0 172889-27-9
          1105 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L102
FILE 'HCAPLUS'
          1137 PYRAZOLOPYRIMIDINE#
          1187 PP1
           274 PP2
            20 172889-26-8
            10 172889-27-9
          2532 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L103
FILE 'NTIS'
             1 PYRAZOLOPYRIMIDINE#
            15 PP1
             2 PP2
             0 172889-26-8
             0 172889-27-9
            18 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L104
FILE 'ESBIOBASE'
             9 PYRAZOLOPYRIMIDINE#
           694 PP1
           109 PP2
             0 172889
         44754 26
        237842 8
             0 172889-26-8
                 (172889(W)26(W)8)
             0 172889
         42591 27
        177442 9
             0 172889-27-9
                 (172889(W)27(W)9)
           784 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L105
FILE 'BIOTECHNO'
            15 PYRAZOLOPYRIMIDINE#
           571 PP1
            76 PP2
             0 172889-26-8
             0 172889-27-9
           645 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L106
FILE 'WPIDS'
            71 PYRAZOLOPYRIMIDINE#
           140 PP1
            69 PP2
            0 172889
        300469 26
       1687338 8
             0 172889-26-8
                 (172889 (W) 26 (W) 8)
             0 172889
```

```
186349 27
      1310082 9
           0 172889-27-9
               (172889(W)27(W)9)
          225 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L107
TOTAL FOR ALL FILES
         9514 (PYRAZOLOPYRIMIDINE# OR PP1 OR PP2 OR 172889-26-8 OR 172889-27-9
L108
=> s 196 and 1108
FILE 'MEDLINE'
     252 L85 AND L97
L109
FILE 'SCISEARCH'
L110
         234 L86 AND L98
FILE 'LIFESCI'
L111 83 L87 AND L99
FILE 'BIOTECHDS'
L112 0 L88 AND L100
FILE 'BIOSIS'
L113 307 L89 AND L101
FILE 'EMBASE'
         217 L90 AND L102
FILE 'HCAPLUS'
         253 L91 AND L103
L115
FILE 'NTIS'
           1 L92 AND L104
L116
FILE 'ESBIOBASE'
        195 L93 AND L105
L117
FILE 'BIOTECHNO'
     135 L94 AND L106
L118
FILE 'WPIDS'
           3 L95 AND L107
L119
TOTAL FOR ALL FILES
     1680 L96 AND L108
L120
=> s 196(5a)1108
FILE 'MEDLINE'
L121 236 L85(5A)L97
FILE 'SCISEARCH'
L122 213 L86(5A)L98
FILE 'LIFESCI'
L123 78 L87(5A)L99
```

FILE 'BIOTECHDS'

L124 0 L88 (5A) L100

FILE 'BIOSIS'

L125 284 L89(5A)L101

FILE 'EMBASE'

L126 194 L90(5A)L102

FILE 'HCAPLUS'

L127 231 L91(5A)L103

FILE 'NTIS'

L128 1 L92(5A)L104

FILE 'ESBIOBASE'

L129 179 L93(5A)L105

FILE 'BIOTECHNO'

L130 128 L94(5A)L106

FILE 'WPIDS'

L131 1 L95(5A)L107

TOTAL FOR ALL FILES

L132 1545 L96(5A) L108

=> s 124 and 1108

FILE 'MEDLINE'

L133 0 L13 AND L97

FILE 'SCISEARCH'

L134 0 L14 AND L98

FILE 'LIFESCI'

L135 0 L15 AND L99

FILE 'BIOTECHDS'

L136 0 L16 AND L100

FILE 'BIOSIS'

L137 1 L17 AND L101

FILE 'EMBASE'

L138 0 L18 AND L102

FILE 'HCAPLUS'

L139 1 L19 AND L103

FILE 'NTIS'

L140 0 L20 AND L104

FILE 'ESBIOBASE'

L141 0 L21 AND L105

FILE 'BIOTECHNO'

L142 0 L22 AND L106

FILE 'WPIDS'

L143 0 L23 AND L107

TOTAL FOR ALL FILES L144 2 L24 AND L108 => s 148 and 1108 FILE 'MEDLINE' 10 L37 AND L97 1.145FILE 'SCISEARCH' 14 L38 AND L98 L146 FILE 'LIFESCI' 2 L39 AND L99 L147 FILE 'BIOTECHDS' 0 L40 AND L100 L148 FILE 'BIOSIS' 14 L41 AND L101 L149 FILE 'EMBASE' L150 8 L42 AND L102 FILE 'HCAPLUS' L151 15 L43 AND L103 FILE 'NTIS' L152 0 L44 AND L104 FILE 'ESBIOBASE' 8 L45 AND L105 L153 FILE 'BIOTECHNO' 4 L46 AND L106 L154 FILE 'WPIDS' 1 L47 AND L107 L155 TOTAL FOR ALL FILES 76 L48 AND L108 L156 => s (136 or 184 or 1144 or 1156) not 2001-2002/py FILE 'MEDLINE' 501323 2001-2002/PY 33 (L25 OR L73 OR L133 OR L145) NOT 2001-2002/PY L157 FILE 'SCISEARCH' 969262 2001-2002/PY 39 (L26 OR L74 OR L134 OR L146) NOT 2001-2002/PY L158 FILE 'LIFESCI' 64358 2001-2002/PY 15 (L27 OR L75 OR L135 OR L147) NOT 2001-2002/PY FILE 'BIOTECHDS' 11462 2001-2002/PY

0 (L28 OR L76 OR L136 OR L148) NOT 2001-2002/PY

FILE 'BIOSIS'

461214 2001-2002/PY 38 (L29 OR L77 OR L137 OR L149) NOT 2001-2002/PY L161 FILE 'EMBASE' 433129 2001-2002/PY 35 (L30 OR L78 OR L138 OR L150) NOT 2001-2002/PY 1.162FILE 'HCAPLUS' 1027851 2001-2002/PY 32 (L31 OR L79 OR L139 OR L151) NOT 2001-2002/PY L163 FILE 'NTIS' 0 2001-2002/PY 0 (L32 OR L80 OR L140 OR L152) NOT 2001-2002/PY L164 FILE 'ESBIOBASE' 271702 2001-2002/PY 27 (L33 OR L81 OR L141 OR L153) NOT 2001-2002/PY L165 FILE 'BIOTECHNO' 114906 2001-2002/PY 25 (L34 OR L82 OR L142 OR L154) NOT 2001-2002/PY L166 FILE 'WPIDS' 889018 2001-2002/PY 0 (L35 OR L83 OR L143 OR L155) NOT 2001-2002/PY L167 TOTAL FOR ALL FILES 244 (L36 OR L84 OR L144 OR L156) NOT 2001-2002/PY => dup rem 1168 PROCESSING COMPLETED FOR L168 53 DUP REM L168 (191 DUPLICATES REMOVED) L169 => d tot MEDLINE DUPLICATE 1 L169 ANSWER 1 OF 53 Collagen, convulxin, and thrombin stimulate aggregation-independent ΤТ tyrosine phosphorylation of CD31 in platelets. Evidence for the involvement of Src family kinases. JOURNAL OF BIOLOGICAL CHEMISTRY, (2000 Sep 1) 275 (35) 27339-47. SO Journal code: HIV; 2985121R. ISSN: 0021-9258. Cicmil M; Thomas J M; Sage T; Barry F A; Leduc M; Bon C; Gibbins J M ΑU 2000472698 MEDLINE ΑN L169 ANSWER 2 OF 53 DUPLICATE 2 MEDLINE Norepinephrine induces vascular endothelial growth TΙ factor gene expression in brown adipocytes through a beta -adrenoreceptor/cAMP/protein kinase A pathway involving Src but independently of Erk1/2. JOURNAL OF BIOLOGICAL CHEMISTRY, (2000 May 5) 275 (18) 13802-11. SO Journal code: HIV; 2985121R. ISSN: 0021-9258. Fredriksson J M; Lindquist J M; Bronnikov G E; Nedergaard J ΑU 2000250944 MEDLINE ΑN DUPLICATE 3 L169 ANSWER 3 OF 53 MEDLINE Oncogenes and tumor angiogenesis: the HPV-16 E6 oncoprotein activates the

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- L169 ANSWER 6 OF 53 MEDLINE

- DUPLICATE 6
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PATENT NO. KIND DATE APPLICATION NO. DATE

PI WO 9816638 A1 19980423 WO 1996-US16495 19961016 W: AU, CA, JP, MX

RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE AU 9674324 Al 19980511 AU 1996-74324 19961016

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L121 ANSWER 7 OF 53 MEDLINE

DUPLICATE 7

Vascular endothelial growth factor (VEGF) appears to be a critical cytokine modulating the growth and spread of Kaposi's sarcoma (KS). Furthermore, infection with the KS herpes virus results in up-regulation of **VEGF** and triggering of **VEGF** receptor activation. The molecular mechanisms regulating such cytokine-driven proliferation of KS cells are not well characterized. We investigated the role of **Src**-related tyrosine kinases in VEGF-mediated signaling in model KS 38 tumor cells. VEGF stimulation specifically activated c-Src kinase activity but not that of other related Src kinases such as Lyn, Fyn, or Hck in KS cells. Pyrazolopyrimidine, a selective inhibitor of Src family tyrosine kinases, significantly blocked the VEGF-induced growth of KS cells. Further studies using mutants of c-Src kinase revealed that Src mediates mitogen-activated protein kinase activation induced by VEGF. We also observed that **VEGF** stimulation resulted in increased tyrosine phosphorylation of the focal adhesion components paxillin and pl30cas. Furthermore, **VEGF** induction enhanced the complex formation between **Src** kinase and paxillin. Src kinase appears to play an important functional role in VEGF-induced signaling in KS cells and may act to link pathways from the VEGF receptor to mitogen-activated protein kinase and cytoskeletal components, thereby effecting tumor proliferation and migration.

L121 ANSWER 11 OF 53 SCISEARCH COPYRIGHT 2002 ISI (R) DUPLICATE 10

DUPLICATE 12 L121 ANSWER 15 OF 53 MEDLINE Vascular endothelial growth factor (VEGF) is a potent endothelial AB cell-specific mitogen that promotes angiogenesis, vascular hyperpermeability, and vasodilation by autocrine mechanisms involving nitric oxide (NO) and prostacyclin (PGI(2)) production. These experiments used immunoprecipitation and immunoassay procedures to characterize the signaling pathways by which VEGF induces NO and PGI(2) formation in cultured endothelial cells. The data showed that VEGF stimulates complex formation of the flk-1/kinase-insert domain-containing receptor (KDR) **VEGF** receptor with c-Src and that Src activation is required for **VEGF** induction of phospholipase C gammal activation and inositol 1,4,5-trisphosphate formation. Reporter cell assays showed that VEGF promotes a approximately 50-fold increase in NO formation, which peaks at 5-20 min. This effect is mediated by a signaling cascade initiated by flk-1/KDR activation of c-Src, leading to phospholipase C gammal activation, inositol 1,4,5-trisphosphate formation, release of [Ca(2+)](i) and nitric oxide synthase activation. Immunoassays of VEGF-induced 6-keto prostaglandin F(lalpha) formation as an indicator of PGI(2) production revealed a 3-4-fold increase that peaked at 45-60 min. The PGI(2) signaling pathway follows the NO pathway through release of [Ca(2+)](i), but diverges prior to NOS activation and also requires activation of mitogen-activated protein kinase. These results suggest that NO and PGI(2) function in parallel in mediating the effects of VEGF.

L121 ANSWER 20 OF 53 MEDLINE

DUPLICATE 16

- Angiogenesis is a prerequisite for solid tumor growth. Glioblastoma multiforme, the most common malignant brain tumor, is characterized by extensive vascular proliferation. We previously showed that transgenic mice expressing a GFAP-v-src fusion gene in astrocytes develop low-grade astrocytomas that progressively evolve into hypervascularized glioblastomas. Here, we examined whether tumor progression triggers angiogenetic signals. We found abundant transcription of vascular endothelial growth factor (VEGF) in neoplastic astrocytes at surprisingly early stages of tumorigenesis. VEGF and v-src expression patterns were not identical, suggesting that VEGF activation was not only dependent on V-src. Late-stage gliomas showed perinecrotic VEGF up-regulation similarly to human glioblastoma. Expression patterns of the endothelial angiogenic receptors flt-1, flk-1, tie-1, and tie-2 were similar to those described in human gliomas, but flt-1 was expressed also in neoplastic astrocytes, suggesting an autocrine role in tumor growth. In crossbreeding experiments, hemizygous ablation of the tumor suppressor genes Rb and p53 had no significant effect on the expression of VEGF, flt-1, flk-1, tie-1, and tie-2. Therefore, expression of angiogenic signals is an early event during progression of GFAP-v-src tumors and precedes hypervascularization. Given the close similarities in the progression pattern between GFAP-v-src and human gliomas, the present results suggest that these mice may provide a useful tool for antiangiogenic therapy research.
- L121 ANSWER 30 OF 53 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.DUPLICATE 21 Tyrosine kinases, first described as oncogenes, have been shown to play a AΒ role in normal cellular processes. Aberrations in tyrosine kinase activity lead to disease states. For fifteen years it has been postulated that the inhibition of tyrosine kinases may have therapeutic utility and the design and testing of inhibitors have been major focuses of research and development in both academic institutions and pharmaceutical companies. While early research focused on developing chemical entities that mimic phosphotyrosine, later research has focused on developing competitive adenosine triphosphate (ATP) inhibitors with various levels of selectivity on kinase targets. This review focuses on a discussion of tyrosine kinases thought to be important in disease, including platelet-derived growth factor (PDGF), fibroblast growth factor (FGF), vascular endothelial cell growth factor (VEGF), epidermal growth factor (EGF) receptors, HER-2 and Src. In addition, the classes of inhibitors designed to affect these targets and that have overcome research and development challenges and entered clinical trials are discussed. These include isoxazole, quinazoline, substituted pyrimidines and indolinone compounds, all of which are in clinical trials or near clinical development by SUGEN, Zeneca, Novartis, Pfizer and Parke-Davis. A summary of the chemistry and activity of these agents is provided.

- Hemodynamic abnormalities have been implicated in the pathogenesis of the AΒ increased glomerular permeability to protein of diabetic and other glomerulopathies. Vascular permeability factor (VPF) is one of the most powerful promoters of vascular permeability. We studied the effect of stretch on VPF production by human mesangial cells and the intracellular signaling pathways involved. The application of mechanical stretch (elongation 10%) for 6 h induced a 2.4-fold increase over control in the VPF mRNA level (P < 0.05). There was a corresponding 3-fold increase in VPF protein level by 12 h (P < 0. 001), returning to the baseline by 24 h. Stretch-induced VPF secretion was partially prevented both by the protein kinase C (PKC) inhibitor H7 (50 microM: $72^{\frac{1}{2}}$ inhibition, P < 0.05) and by pretreatment with phorbol ester (phorbol-12-myristate-13 acetate $10\,(-)\,7$ M: 77* inhibition, P < 0.05). A variety of protein tyrosine kinase (PTK) inhibitors, genistein (20 microg/ml), herbimycin \bar{A} (3.4 microM), and a specific pp60(src) peptide inhibitor (21 microM) also significantly reduced, but did not entirely prevent, stretch-induced **VPF** protein secretion (respectively 63*, 80*, and 75% inhibition; P < 0.05 for all). The combination of both PKC and PTK inhibition completely abolished the VPF response to mechanical stretch (100% inhibition, P < 0.05). Stretch induces VPF gene expression and protein secretion in human mesangial cells via PKC- and PTK-dependent mechanisms.
- L121 ANSWER 39 OF 53 SCISEARCH COPYRIGHT 2002 ISI (R) DUPLICATE 27 Balloon angioplasty disrupts the protective endothelial lining of the AΒ arterial wall, rendering arteries susceptible to thrombosis and intimal thickening. We show here that Vascular endothelial growth factor (VEGF), an endothelial cell mitogen, is upregulated in medial smooth muscle cells of the arterial wall in response to balloon injury. Both protein kinase C (PKC) and tyrosine kinase pp60(src) mediate augmented VEGF expression. In contrast, nitric oxide (NO) donors inhibit PKC-induced **VEGF** upregulation by interfering with binding of the transcription factor activator protein-1 (AP-1) to the VEGF promoter. Inhibition of VEGF promoter activation suggests that NO secreted by a restored endothelium functions as the negative feedback mechanism that downregulates VEGF expression to basal levels. Administration of a neutralizing VEGF antibody impaired reendothelialization following balloon injury performed in vivo. These findings establish a reciprocal relation between VEGF and NO in the endogenous regulation of endothelial integrity following arterial injury.
- DUPLICATE 35 L121 ANSWER 52 OF 53 MEDLINE Vascular endothelial growth factor (VEGF) is a homodimeric peptide growth AB factor which binds to two structurally related tyrosine kinase receptors denoted Flt1 and KDR. In order to compare the signal transduction via these two receptors, the human Flt1 and KDR proteins were stably expressed in porcine aortic endothelial cells. Binding analyses using 125I-VEGF revealed Kd values of 16 pM for Flt1 and 760 pM for KDR. Cultured human umbilical vein endothelial (HUVE) cells were found to express two distinct populations of binding sites with affinities similar to those for Flt1 and KDR, respectively. The KDR expressing cells showed striking changes in cell morphology, actin reorganization and membrane ruffling, chemotaxis and mitogenicity upon VEGF stimulation, whereas Fltl expressing cells lacked such responses. KDR was found to undergo ligand-induced autophosphorylation in intact cells, and both Flt1 and KDR were phosphorylated in vitro in response to VEGF, however, KDR much more efficiently than Flt1. Neither the receptor-associated activity of phosphatidylinositol 3'-kinase nor tyrosine phosphorylation of phospholipase C-gamma were affected by stimulation of Fltl or KDR expressing cells, and phosphorylation of GTPase activating protein was

only slightly increased. Members of the **Src** family such as Fyn and **Yes** showed an increased level of phosphorylation upon **VEGF** stimulation of cells expressing Flt1 but not in cells expressing KDR. The maximal responses in KDR expressing porcine aortic endothelial cells were obtained at higher VEGF concentrations as compared to HUVE cells, i.e. in the presence of Flt1. This difference could possibly be explained by the formation of heterodimeric complexes between KDR and Flt1, or other molecules, in HUVE cells.

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